From the INTERNATIONAL SEARCHING AUTHORITY

To:

BARKER BRETTELL 10-12 Priests Bridge London SW15 5JE UNITED KINGDOM

BARKER PRETTELL LONDON

PCT

NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION

(PCT Rule 44.1)

Date of mailing (day/month/year) 10/06/2003 Applicant's or agent's file reference JPP183 FOR FURTHER ACTION See paragraphs 1 and 4 below International application No. International filing date PCT/GB 03/00966 (day/month/year) 07/03/2003 Applicant ROYAL HOLLOWAY UNIVERSITY OF LONDON

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1	- X	The appl	icant is hereby r	notified that the International Search Report has been established and is transmitted herewith
		rungor	amendments a	and Statement under Article 10.
		тпе аррі	icant is entitled,	if he so wishes, to amend the claims of the International Application (see Rule 46):
		When?	The time limit fo International Se	or filing such amendments is normally 2 months from the date of transmittal of the earch Report; however, for more details, see the notes on the accompanying sheet.
		Where?	Directly to the	International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41–22) 740.14.35
		For more	detailed instru	uctions, see the notes on the accompanying sheet.
2		The appli Article 17	cant is hereby no (2)(a) to that effo	otified that no International Search Report will be established and that the declaration under ect is transmitted herewith.
3.		With rega	ard to the prote	st against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that:
	,	the pappl	protest together icant's request to	with the decision thereon has been transmitted to the International Bureau together with the offices.
	l	no d	ecision has beer	n made yet on the protest; the applicant will be notified as soon as a decision is made.
4.	Furth	er action(s): The applic	cant is reminded of the following:
	com	ity claim, in pletion of	must reach the I the technical pre	e priority date, the international application will be published by the International Bureau. d or postpone publication, a notice of withdrawal of the international application, or of the nternational Bureau as provided in Rules 90 <i>bis</i> .1 and 90 <i>bis</i> .3, respectively, before the parations for international publication.
			•	ity date, a demand for international preliminary examination must be filed if the applicant nto the national phase until 30 months from the priority date (in some Offices even later).
	Within before	20 month	is from the priori	ity date, the applicant must perform the prescribed acts for entry into the national phase

before all designated Offices which have not been elected in the demand or in a later election within 19 months from the

Name and mailing address of the International Searching Authority



European Patent Office, P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,

priority date or could not be elected because they are not bound by Chapter II.

Fax: (+31-70) 340-3016

Authorized officer

Sabine Oblinger

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international publication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been is filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

Notes to Form PCT/ISA/220 (first sheet) (January 1994)

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers; claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- (Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims):
 Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added. or
 Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged.
- [Where various kinds of amendments are made]:

 "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

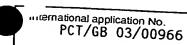
For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.



INTERNATIONAL SEARCH REPORT

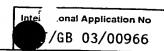
(PCT Article 18 and Rules 43 and 44)

JPP183	eference FOR FURTHEI ACTION	R see Notification of (Form PCT/ISA/2	of Transmittal of International Search Report 220) as well as, where applicable, item 5 below
International application No	o. International filing date ((day/month/year)	(Earliest) Priority Date (day/month/year)
PCT/GB 03/00966	07/03/2	2003	07/03/2002
Applicant			07/03/2002
ROYAL HOLLOWAY U	NIVERSITY OF LONDON		
This International Search according to Article 18. A	Report has been prepared by this Internaticopy is being transmitted to the Internation	ional Searching Auth nal Bureau.	ority and is transmitted to the applicant
This International Search I	Report consists of a total of5 accompanied by a copy of each prior art do	sheets.	report.
1. Basis of the report			
	language, the international search was ca it was filed, unless otherwise indicated und	dor tino nem.	
the interna Authority (i	tional search was carried out on the basis Rule 23.1(b)).	of a translation of the	e international application furnished to this
b. With regard to any	nucleotide and/or amino soid some	e disclosed in the inte	ernational application, the international search
	the basis of the sequence listing: in the international application in written for		.,, s.e mematerial search
	ner with the international application in com		
furnished s	subsequently to this Authority in written form	n m	•
	subsequently to this Authority in computer r		
	ent that the subsequently furnished written al application as filed has been furnished.		
the stateme furnished	ent that the information recorded in comput	ter readable form is i	dentical to the written sequence listing has bee
. X Certain cla	ims were found unsearchable (See Box	1).	
Unity of inv	vention is lacking (see Box II).		
. With regard to the title,			
the text is a	pproved as submitted by the applicant.		
	been established by this Authority to read	as follows:	
RECOMBINANT SP	PORES		
With regard to the abstr	act,		
X the text is ap	proved as submitted by the applicant.		
the text has i	been established, according to Rule 38.2(to the stablished in the date of mailing of this international to the stable in the sta	b), by this Authority a	as it appears in Box III. The applicant may,
The figure of the drawing	gs to be published with the abstract is Figu	ure No.	1
	by the applicant.		None of the figures.
			I I INVITE OF THE TIGHTEE
X because the	applicant failed to suggest a figure.		



	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
	emational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
1	Although claims 30 and 31 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
1 - 1	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
b	Claims Nos.: ecause they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II O	bservations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Interna	ational Searching Authority found multiple inventions in this international application, as follows:
	application, as follows:
1. As	all required additional search fees were timely paid by the applicant, this International Search Report covers all archable claims.
500	archable claims.
	is an interport covers all
	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment
2. As a of a	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment any additional fee.
2. As a of a	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment additional fee.
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2. As a of a	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment additional fee.
2. As a of a	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment additional fee.
2. As a of a 3. As o cove	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment invalue additional fee. Only some of the required additional search fees were timely paid by the applicant, this International Search Report ers only those claims for which fees were paid, specifically claims Nos.:
2. As a of a 3. As o cove	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment additional fee.
2. As a of a 3. As o cove	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment invalue additional fee. Only some of the required additional search fees were timely paid by the applicant, this International Search Report ers only those claims for which fees were paid, specifically claims Nos.:
2. As a of a 3. As o cove	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment invalue additional fee. Only some of the required additional search fees were timely paid by the applicant, this International Search Report ers only those claims for which fees were paid, specifically claims Nos.:
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2. As a of a o	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment any additional fee. Some of the required additional search fees were timely paid by the applicant, this International Search Report sers only those claims for which fees were paid, specifically claims Nos.: Sequired additional search fees were timely paid by the applicant. Consequently, this International Search Report is covered to the invention first mentioned in the claims; it is covered by claims Nos.:
2. As a of a 3. As o cove	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment any additional fee. Some of the required additional search fees were timely paid by the applicant, this International Search Report sers only those claims for which fees were paid, specifically claims Nos.: Sequired additional search fees were timely paid by the applicant. Consequently, this International Search Report is covered to the invention first mentioned in the claims; it is covered by claims Nos.:
2. As a of a o	all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment any additional fee. Only some of the required additional search fees were timely paid by the applicant, this International Search Report are only those claims for which fees were paid, specifically claims Nos.: Sequired additional search fees were timely paid by the applicant. Consequently, this International Search Report is cited to the invention first mentioned in the claims; it is covered by claims Nos.:

IN' RNATIONAL SEARCH REPORT



A. CLASSIFICATION OF SUBJECT HEATTER IPC 7 C12N3/00 C12N15/00

A61K39/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) C12N A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, WPI Data, PAJ, MEDLINE, EMBASE

Catana		
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 02 00232 A (MAXYGEN INC ;GOLDMAN STANLEY (US); WHALEN ROBERT G (US); LATHROP S) 3 January 2002 (2002-01-03)	1-8, 10-12, 17,18,
r	in particular, examples and claims. the whole document	23-31 11
(WO 01 94599 A (KATAGIHALLIMATH NAINESH ;MUKHERJEE KAKOLI (IN); SMITTSKYDDSINSTITU) 13 December 2001 (2001-12-13)	1-13, 15-20, 23-31
	the whole document	11
	-/	
X Furthe	r documents are listed in the continuation of box C. X Patent family members	are listed in annex.

X Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents:	
'A' document defining the general state of the art which is not considered to be of particular relevance 'E' earlier document but published on or after the international filing date 'L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) 'O' document referring to an oral disclosure, use, exhibition or other means 'P' document published prior to the international filing date but later than the priority date claimed	 'T' later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention 'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone 'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. '&' document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
26 May 2003	10/06/2003
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk	Authorized officer
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Rojo Romeo, E

1

IN TRNATIONAL SEARCH REPORT

lnte'	ional Application No	
	/GB 03/00966	

C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	T/GB 03/00966
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y		rielevant to claim No.
	ROBINSON K ET AL: "ORAL VACCINATION OF MICE AGAINST TETANUS WITH RECOMBINANT LACTOCOCCUS LACTIS" NATURE BIOTECHNOLOGY, NATURE PUBLISHING, US, vol. 15, no. 7, 15 July 1997 (1997-07-15), pages 653-657, XP001093957 ISSN: 1087-0156 the whole document	
	MINTON NIGEL P ET AL: "Chemotherapeutic tumour targeting using clostridial spores." FEMS MICROBIOLOGY REVIEWS, vol. 17, no. 3, 1995, pages 357-364, XP002242524 ISSN: 0168-6445 the whole document	21,22

IN PRNATIONAL SEARCH REPORT

Information on patent family members

Inter ional	Application No
T/GB	03/00966
mih.	

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 0200232	Α	03-01-2002	AU EP WO US	7300901 A 1299115 A2 0200232 A2 2002150594 A1	08-01-2002 09-04-2003 03-01-2002 17-10-2002
WO 0194599	A	13-12-2001	AU EP WO	6448801 A 1292686 A1 0194599 A1	17-12-2001 19-03-2003 13-12-2001

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